

Reisolation of plaques for cDNA cloning, sequencing and reconstruction of previously designed replicons and helpers

FIG. 1

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ATTGACGGCGTAGTACACACTATTGAATCAAACAGCCGACCAATTGCACTACCATCACAATGGAGAAGCCAGTAG TAAACGTAGACGTAGACCCCCAGAGTCCGTTTGTCGTGCAACTGCAAAAAAGCTTCCCGCAATTTGAGGTAGTAG CACAGCAGGTCACTCCAAATGACCATGCTAATGCCAGAGCATTTTCGCATCTGGCCAGTAAACTAATCGAGCTGG AGGTTCCTACCACAGCGACGATCTTGGACATAGGCAGCGCACCGGCTCGTAGAATGTTTTCCGAGCACCAGTATC ATTGTGTCTGCCCCATGCGTAGTCCAGAAGACCCGGACCGCATGATGAAATATGCCAGTAAACTGGCGGAAAAAG CGTGCAAGATTACAAACAAGAACTTGCATGAGAAGATTAAGGATCTCCGGACCGTACTTGATACGCCGGATGCTG AAACACCATCGCTCTGCTTTCACAACGATGTTACCTGCAACATGCGTGCCGAATATTCCGTCATGCAGGACGTGT ATATCAACGCTCCCGGAACTATCTATCATCAGGCTATGAAAGGCGTGCGGACCCTGTACTGGATTGGCTTCGACA CCACCCAGTTCATGTTCTCGGCTATGGCAGGTTCGTACCCTGCGTACAACACCAACTGGGCCGACGAGAAAGTCC TTGAAGCGCGTAACATCGGACTTTGCAGCACAAAGCTGAGTGAAGGTAGGACAGGAAAATTGTCGATAATGAGGA AGAAGGAGTTGAAGCCCGGGTCGCGGGTTTATTTCTCCGTAGGATCGACACTTTATCCAGAACACAGAGCCAGCT TGCAGAGCTGGCATCTTCCATCGGTGTTCCACTTGAATGGAAAGCAGTCGTACACTTGCCGCTGTGATACAGTGG TGAGTTGCGAAGGCTACGTAGTGAAGAAAATCACCATCAGTCCCGGGATCACGGGAGAAACCGTGGGATACGCGG TTACACACAATAGCGAGGGCTTCTTGCTATGCAAAGTTACTGACACAGTAAAAGGAGAACGGGTATCGTTCCCTG TGTGCACGTACATCCCGGCCACCATATGCGATCAGATGACTGGTATAATGGCCACGGATATATCACCTGACGATG CACAAAAACTTCTGGTTGGGCTCAACCAGCGAATTGTCATTAACGGTAGGACTAACAGGAACACCAACACCATGC AAAATTACCTTCTGCCGATCA'IAGCACAAGGGTTCAGCAAATGGGCTAAGGAGCGCAAGGATGATCTTGATAACG AGAAATGCTGGGTACTAGAGAACGCAAGCTTACGTACGGCTGCTTGTGGGCCGTTTCGCACTAAGAAAGTACATT CGTTTTATCGCCCACCTGGAACGCAGACCATCGTAAAAGTCCCAGCCTCTTTTAGCGCTTTTCCCATGTCGTCCG TATGGACGACCTCTTTGCCCATGTCGCTGAGGCAGAAATTGAAACTGGCATTGCAACCAAAGAAGAAGGAAAAAAAC TGCTGCAGGTCTCGGAGGAATTAGTCATGGAGGCCAAGGCTGCTTTTGAGGATGCTCAGGAGGAAGCCAGAGCGG AGAAGCTCCGAGAAGCACTTCCACCATTAGTGGCAGACAAAGGCATCGAGGCAGCCGCAGAAGTTGTCTGCGAAG TGGAGGGGCTCCAGGCGGACATCGGAGCAGCATTAGTTGAAACCCCGCGCGCTCACGTAAGGATAATACCTCAAG CAAATGACCGTATGATCGGACAGTATATCGTTGTCTCGCCAAACTCTGTGCTGAAGAATGCCAAACTCGCACCAG CGCACCCGCTAGCAGATCAGGTTAAGATCATAACACACTCCGGAAGATCAGGAAGGTACGCGGTCGAACCATACG **AAGAGGAGCAGTACAAGGTTACAAAGGCAGAGCTTGCAGAAACAGAGTACGTGTTTGACGTGGACAAGAAGCGTT** TGGAGGGACTGAAGACCCGACCTGCGGTCCCGTACAAGGTCGAAACAATAGGAGTGATAGGCACACCGGGGTCGG AAATTGAGGCCGACGTGCTAAGACTGAGGGGTATGCAGATTACGTCGAAGACAGTAGATTCGGTTATGCTCAACG GATGCCACAAAGCCGTAGAAGTGCTGTACGTTGACGAAGCGTTCGCGTGCCACGCAGGAGCACTACTTGCCTTGA TTGCTATCGTCAGGCCCCGCAAGAAGGTAGTACTATGCGGAGACCCCATGCAATGCGGATTCTTCAACATGATGC **AACTAAAGGTACATTTCAATCACCCTGAAAAAGACATATGCACCAAGACATTCTACAAGTATATCTCCCGGCGTT** GCACACAGCCAGTTACAGCTATTGTATCGACACTGCATTACGATGGAAAGATGAAAACCACGAACCCGTGCAAGA GGGTTAAGCAATTGCAAATCGACTATCCCGGACATGAAGTAATGACAGCCGCGGCCTCACAAGGGCTAACCAGAA AAGGAGTGTATGCCGTCCGGCAAAAAGTCAATGAAAACCCACTGTACGCGATCACATCAGAGCATGTGAACGTGT TGCTCACCCGCACTGAGGACAGGCTAGTGTGGAAAACCTTGCAGGGCGACCCATGGATTAAGCAGCTCACTAACA TACCTAAAGGAAACTTTCAGGCTACTATAGAGGACTGGGAAGCTGAACACAAGGGAATAATTGCTGCAATAAACA GCCCCACTCCCGTGCCAATCCGTTCAGCTGCAAGACCAACGTTTGCTGGGCGAAAGCATTGGAACCGATACTAG CCACGCCGGTATCGTACTTACCGGTTGCCAGTGGAGCGAACTGTTCCCACAGTTTGCGGATGACAAACCACATT AGAGCATCCCACTAACGTACCATCCCGCCGATTCAGCGAGGCCGGTAGCTCATTGGGACAACAGCCCAGGAACCC GCACACACTTGATTTGCAGACGGGGAGAACCAGAGTTATCTCTGCACAGCATAACCTGGTCCCGGTGAACCGCA ATCTTCCTCACGCCTTAGTCCCCGAGTACAAGGAGAAGCAACCCGGCCCGGTCGAAAAATTCTTGAACCAGTTCA AACACCACTCAGTACTTGTGGTATCAGAGGAAAAAATTGAAGCTCCCCGTAAGAGAATCGAATGGATCGCCCCGA TTGGCATAGCCGGTGCAGATAAGAACTACAACCTGGCTTTCGGGTTTCCGCCGCAGGCACGGTACGACCTGGTGT TCATCAACATTGGAACTAAATACAGAAACCACCACTTTCAGCAGTGCGAAGACCATGCGGCGACCTTAAAAACCC TTTCGCGTTCGGCCCTGAATTGCCTTAACCCAGGAGGCACCCTCGTGGTGAAGTCCTATGGCTACGCCGACCGCA ACAGTGAGGACGTAGTCACCGCTCTTGCCAGAAAGTTTGTCAGGGTGTCTGCAGCGAGACCAGATTGTGTCTCAA

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ATTGCGTGATTTCGTCCGTGTATGAGGGTACAAGAGATGGAGTTGGAGCCGCGCCGTCATACCGCACCAAAAGGG AGAATATTGCTGACTGTCAAGAGGAAGCAGTTGTCAACGCAGCCAATCCGCTGGGTAGACCAGGCGAAGGAGTCT GCCGTGCCATCTATAAACGTTGGCCGACCAGTTTTACCGATTCAGCCACGGAGACAGGCACCGCAAGAATGACTG TGTGCCTAGGAAAGAAAGTGATCCACGCGGTCGGCCCTGATTTCCGGAAGCACCCAGAAGCAGAAGCCTTGAAAT TGCTACAAAACGCCTACCATGCAGTGGCAGACTTAGTAAATGAACATAACATCAAGTCTGTCGCCATTCCACTGC TATCTACAGGCATTTACGCAGCCGGAAAAGACCGCCTTGAAGTATCACTTAACTGCTTGACAACCGCGCTAGACA GCTTGAAGGGAAGAAAGGGATTCAGTACTACAAAAGGAAAATTGTATTCGTACTTCGAAGGCACCAAATTCCATC AAGCAGCAAAAGACATGGCGGAGATAAAGGTCCTGTTCCCTAATGACCAGGAAAGTAATGAACAACTGTGTGCCT AAACGTTGCCGTGCCTTTGCATGTATGCCATGACGCCAGAAAGGGTCCACAGACTTAGAAGCAATAACGTCAAAG AAGTTACAGTATGCTCCTCCACCCCCCTTCCTAAGCACAAAATTAAGAATGTTCAGAAGGTTCAGTGCACGAAAG CTCCTCCTGCACAGGCCGAGGAGGCCCCCGAAGTTGTAGCGACACCGTCACCATCTACAGCTGATAACACCTCGC TTGATGTCACAGACATCTCACTGGATATGGATGACAGTAGCGAAGGCTCACTTTTTTCGAGCTTTAGCGGATCGG ACAACTCTATTACTAGTATGGACAGTTGGTCGTCAGGACCTAGTTCACTAGAGATAGTAGACCGAAGGCAGGTGG TGGTGGCTGACGTTCATGCCGTCCAAGAGCCTGCCCCTATTCCACCGCCAAGGCTAAAGAAGATGGCCCGCCTGG CAGCGGCTAGAAAAGAGCCCACTCCACCGGCAAGCAATAGCTCTGAGTCCCTCCACCTCTTTTTGGTGGGGTAT CCATGTCCCTCGGATCAATTTTCGACGGAGAGACGGCCCGCCAGGCAGCGGTACAACCCCTGGCAACAGGCCCCA CGGATGTGCCTATGTCTTTCGGATCGTTTTCCGACGGAGAGATTGATGAGCTGAGCCGCAGAGTAACTGAGTCCG AACCCGTCCTGTTTGGATCATTTGAACCGGGCGAAGTGAACTCAATTATATCGTCCCGATCAGCCGTATCTTTTC CACTACGCAAGCAGAGCGTAGACGCAGGAGCAGGAGGACTGAATACTGACTAACCGGGGTAGGTGGGTACATAT TTTCGACGGACACAGGCCCTGGGCACTTGCAAAAGAAGTCCGTTCTGCAGAACCAGCTTACAGAACCGACCTTGG AGCGCAATGTCCTGGAAAGAATTCATGCCCCGGTGCTCGACACGTCGAAAGAGAGAACAACTCAAACTCAGGTACC AGATGATGCCCACCGAAGCCAACAAAAGTAGGTACCAGTCTCGTAAAGTAGAAAATCAGAAAGCCATAACCACTG AGCGACTACTGTCAGGACTACGACTGTATAACTCTGCCACAGATCAGCCAGAATGCTATAAGATCACCTATCCGA TGCATGAGAACTATCCGACAGTAGCATCTTATCAGATTACTGACGAGTACGATGCTTACTTGGATATGGTAGACG GGACAGTCGCCTGCCTGGATACTGCAACCTTCTGCCCCGCTAAGCTTAGAAGTTACCCGAAAAAAACATGAGTATA GAGCCCCGAATATCCGCAGTGCGGTTCCATCAGCGATGCAGAACACGCTACAAAATGTGCTCATTGCCGCAACTA AAAGAAATTGCAACGTCACGCAGATGCGTGAACTGCCAACACTGGACTCAGCGACATTCAATGTCGAATGCTTTC GAAAATATGCATGTAATGACGAGTATTGGGAGGAGTTCGCTCGGAAGCCAATTAGGATTACCACTGAGTTTGTCA CCGCATATGTAGCTAGACTGAAAGGCCCTAAGGCCGCCCCCACTATTTGCAAAGACGTATAATTTGGTCCCATTGC AATTAGTGCGTAGGCTTACGGCCGTCTTGCTTCCAAACATTCACACGCTTTTTGACATGTCGGCGGAGGATTTTG ATGCAATCATAGCAGAACACTTCAAGCAAGGCGACCCGGTACTGGAGACGGATATCGCATCATTCGACAAAAGCC **AATCCGGAATGTTCCTCACACTTTTTGTCAACACAGTTTTGAATGTCGTTATCGCCAGCAGAGAGTACTAGAAGAGC** GGCTTAAAACGTCCAGATGTGCAGCGTTCATTGGCGACGACAACATCATACATGGAGTAGTATCTGACAAAGAAA TGGCTGAGAGGTGCGCCACCTGGCTCAACATGGAGGTTAAGATCATCGACGCAGTCATCGGTGAGAGACCACCTT ACTTCTGCGGCGGATTTATCTTGCAAGATTCGGTTACTTCCACAGCGTGCCGCGTGGCGGACCCcctgaaaaggc tgtttaagttgggtaaaccgctcccagccgacgacgacgagcaagacgagacagaagacgcgctctgctagatgaaa caaaggcgtggtttagagtaggtataacaggcactttagcagtggccgtgacgacccggtatgaggtagacaata ttacacctgtcctactggcattgagaacttttgcccagagcaaaagagcattccaagccatcagaggggaaataa agcatctctacggtggtcctaaatagtcagcatagtacatttcatctgactaatactacaacaccaccaccatga atagaggattetttaaeatgeteggeegeegeeetteeeggeeeeeaetgeeatgtggaggeegeggAGAAGGA GGCAGGCGCCCGATGCCTGCCCGCAACGGCTGGCTTCTCAAATCCAGCAACTGACCACAGCCGTCAGTGCCC TAGTCATTGGACAGGCAACTAGACCTCAACCCCCACGTCCACGCCCACCGCCACAGAAGAAGCAGGCGCCCA AGAGACAGCGCATGGCACTTAAGTTGGAGGCCGACAGATCGTTCGACGTCAAGAACGAGGACGGAGATGTCATCG GGCACGCACTGGCCATGGAAGGAAAGGTAATGAAACCTCTGCACGTGAAAGGAACCATCGACCACCCTGTGCTAT CAAAGCTCAAATTTACCAAGTCGTCAGCATACGACATGGAGTTCGCACAGTTGCCAGTCAACATGAGAAGTGAGG CATTCACCTACACCAGTGAACACCCCGAAGGATTCTATAACTGGCACCACGGAGCGGTGCAGTATAGTGGAGGTA GATTTACCATCCCTCGCGGAGTAGGAGGCAGAGGAGACAGCGGTCGTCCGATCATGGATAACTCCGGTCGGGTTG



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TCGCGATAGTCCTCGGTGGAGCTGATGAAGGAACACGAACTGCCCTTTCGGTCGTCACCTGGAATAGTAAAGGGA AGACAATTAAGACGACCCCGGAAGGGACAGAAGAGTGGTCCGCAGCACCACTGGTCACGGCAATGTGTTTGCTCG GAAATGTGAGCTTCCCATGCGACCGCCCGCCCACATGCTATACCCGCGAACCTTCCAGAGCCCTCGACATCCTTG AAGAGAACGTGAACCATGAGGCCTACGATACCCTGCTCAATGCCATATTGCGGTGCGGATCGTCTGGCAGAAGCA AAAGAAGCGTCACTGACGACTTTACCCTGACCAGCCCCTACTTGGGCACATGCTCGTACTGCCACCATACTGAAC CGTGCTTCAGCCCTGTTAAGATCGAGCAGGTCTGGGACGAAGCGGACGATAACACCATACGCATACAGACTTCCG CCCAGTTTGGATACGACCAAAGCGGAGCAGCAAGCGCAAACAAGTACCGCTACATGTCGCTTAAGCAGGATCACA CCGTTAAAGAAGGCACCATGGATGACATCAAGATTAGCACCTCAGGACCGTGTAGAAGGCTTAGCTACAAAGGAT ACTTTCTCCTCGCAAAATGCCCTCCAGGGGACAGCGTAACGGTTAGCATAGTGAGTAGCAACTCAGCAACGTCAT GTACACTGGCCCGCAAGATAAAACCAAAATTCGTGGGACGGGAAAAATATGATCTACCTCCCGTTCACGGTAAAA AAATTCCTTGCACAGTGTACGACCGTCTGAAAGGAACAACTGCAGGCTACATCACTATGCACAGGCCGGGACCGC ACGCTTATACATCCTACCTGGAAGAATCATCAGGGAAAGTTTACGCAAAGCCGCCATCTGGGAAGAACATTACGT ATGAGTGCAAGTGCGGCGACTACAAGACCAGAACCGTTTCGACCCGCACCGAAATCACTGGTTGCACCGCCATCA AGCAGTGCGTCGCCTATAAGAGCGACCAAACGAAGTGGGTCTTCAACTCACCGGACTTGATCAGACATGACGACC ACACGCCCAAGGGAAATTGCATTTGCCTTTCAAGTTGATCCCGAGTACCTGCATGGTCCCTGTTGCCCACGCGC CGAATGTAATACATGGCTTTAAACACATCAGCCTCCAATTAGATACAGACCACTTGACATTGCTCACCACCAGGA GACTAGGGGCAAACCCGGAACCAACCACTGAATGGATCGTCGGAAAGACGGTCAGAAACTTCACCGTCGACCGAG ATGGCCTGGAATACATATGGGGAAATCATGAGCCAGTGAGGGTCTATGCCCAAGAGTCAGCACCAGGAGACCCTC ACGGATGGCCACACGAAATAGTACAGCATTACTACCATCGCCATCCTGTGTACACCATCTTAGCCGTCGCATCAG CATACGCCCTGGCCCCAAACGCCGTAATCCCAACTTCGCTGGCACTCTTGTGCTGCGTTAGGTCGGCCAATGCTG AAACGTTCACCGAGACCATGAGTTACTTGTGGTCGAACAGTCAGCCGTTCTTCTGGGTCCAGTTGTGCATACCTT CGAAGGTAGACGCCTACGAACATGCGACCACTGTTCCAAATGTGCCACAGATACCGTATAAGGCACTTGTTGAAA ACATTACCTGCAAATTCACCACTGTGGTCCCCTCCCCAAAAATCAAATGCTGCGGCTCCTTGGAATGTCAGCCGG CCGTTCATGCAGACTATACCTGCAAGGTCTTCGGAGGGGTCTACCCCTTTATGTGGGGAGGAGCGCAATGTTTTT GCGACAGTGAGAACAGCCAGATGAGTGAGGCGTACGTCGAACTGTCAGCAGATTGCGCGTCTGACCACGCGCAGG CGATTAAGGTGCACACTGCCGCGATGAAAGTAGGACTGCGTATAGTGTACGGGAACACTACCAGTTTCCTAGATG TGTACGTGAACGGAGTCACACCAGGAACGTCTAAAGACTTGAAAGTCATAGCTGGACCAATTTCAGCATCGTTTA CGCCATTCGATCATAAGGTCGTTATCCATCGCGGCCTGGTGTACAACTATGACTTCCCGGAATATGGAGCGATGA AACCAGGAGCGTTTGGAGACATTCAAGCTACCTCCTTGACTAGCAAGGATCTCATCGCCAGCACAGACATTAGGC TACTCAAGCCTTCCGCCAAGAACGTGCATGTCCCGTACACGCAGGCCGCATCAGGATTTGAGATGTGGAAAAACA ACTCAGGCCGCCCACTGCAGGAAACCGCACCTTTCGGGTGTAAGATTGCAGTAAATCCGCTCCGAGCGGTGGACT GTTCATACGGGAACATTCCCATTTCTATTGACATCCCGAACGCTGCCTTTATCAGGACATCAGATGCACCACTGG TCTCAACAGTCAAATGTGAAGTCAGTGAGTGCACTTATTCAGCAGACTTCGGCGGGATGGCCACCCTGCAGTATG TATCCGACCGCGAAGGTCAATGCCCCGTACATTCGCATTCGAGCACAGCAACTCTCCAAGAGTCGACAGTACATG TCCTGGAGAAAGGAGCGGTGACAGTACACTTTAGCACCGCGAGTCCACAGGCGAACTTTATCGTATCGCTGTGTG GGAAGAAGACAACATGCAATGCAGAATGTAAACCACCAGCTGACCATATCGTGAGCACCCCGCACAAAAATGACC AAGAATTTCAAGCCGCCATCTCAAAAACATCATGGAGTTGGCTGTTTGCCCTTTTCGGCGGCGCCCTCGTCGCTAT TAATTATAGGACTTATGATTTTTGCTTGCAGCATGATGCTGACTAGCACACGAAGATGACCGCTACGCCCCAATG ATCCGACCAGCAAAACTCGATGTACTTCCGAGGAACTGATGTGCATAATGCATcaggctggtacattagatcccc gcttaccgcgggcaatatagcaacactaaaaactcgatgtacttccgaggaagcgcagtgcataatgctgcgcag tgttgccacataaccactatattaaccatttatctagcggacgccaaaaactcaatgtatttctgaggaagcgtg gtgcataatgccacgcagcgtctgcataacttttattatttcttttattaatcaacaaaattttgtttttaacat ttc

FIG. 2B-3



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ATTGACGGCGTAGTACACACTATTGAATCAAACAGCCGACCAATTGCACTACCATCACAATGGAGAAGCCAGTAG TAAACGTAGACGTAGACCCCCAGAGTCCGTTTGTCGTGCAACTGCAAAAAAGCTTCCCGCAATTTGAGGTAGTAG CACAGCAGGTCACTCCAAATGACCATGCTAATGCCAGAGCATTTTCGCATCTGGCCAGTAAACTAATCGAGCTGG AGGTTCCTACCACAGCGACGATCTTGGACATAGGCAGCGCACCGGCTCGTAGAATGTTTTCCGAGCACCAGTATC ATTGTGTCTGCCCCATGCGTAGTCCAGAAGACCCGGACCGCATGATGAAATATGCCAGTAAACTGGCGGAAAAAG CGTGCAAGATTACAAACAAGAACTTGCATGAGAAGATTAAGGATCTCCGGACCGTACTTGATACGCCGGATGCTG AAACACCATCGCTCTGCTTTCACAACGATGTTACCTGCAACATGCGTGCCGAATATTCCGTCATGCAGGACGTGT ATATCAACGCTCCCGGAACTATCTATCATCAGGCTATGAAAGGCGTGCGGACCCTGTACTGGATTGGCTTCGACA CCACCCAGTTCATGTTCTCGGCTATGGCAGGTTCGTACCCTGCGTACAACACCAACTGGGCCGACGAGAAAGTCC TTGAAGCGCGTAACATCGGACTTTGCAGCACAAAGCTGAGTGAAGGTAGGACAGGAAAATTGTCGATAATGAGGA **AGAAGGAGTTGAAGCCCGGGTCGCGGGTTTATTTCTCCGTAGGATCGACACTTTATCCAGAACACAGAGCCAGCT** TGCAGAGCTGGCATCTTCCATCGGTGTTCCACTTGAATGGAAAGCAGTCGTACACTTGCCGCTGTGATACAGTGG TGAGTTGCGAAGGCTACGTGGAAGAAAATCACCATCAGTCCCGGGATCACGGGAGAAACCGTGGGATACGCGG TTACACACAATAGCGAGGGCTTCTTGCTATGCAAAGTTACTGACACAGTAAAAGGAGAACGGGTATCGTTCCCTG TGTGCACGTACATCCCGGCCACCATATGCGATCAGATGACTGGTATAATGGCCACGGATATATCACCTGACGATG CACAAAAACTTCTGGTTGGGCTCAACCAGCGAATTGTCATTAACGGTAGGACTAACAGGAACACCAACACCATGC AAAATTACCTTCTGCCGATCATAGCACAAGGGTTCAGCAAATGGGCTAAGGAGCGCAAGGATGATCTTGATAACG AGAAAATGCTGGGTACTAGAGAACGCAAGCTTACGTACGGCTGCTTGTGGGCCGTTTCGCACTAAGAAAGTACATT CGTTTTATCGCCCACCTGGAACGCAGACCATCGTAAAAGTCCCAGCCTCTTTTAGCGCTTTTCCCATGTCGTCCG TATGGACGACCTCTTTGCCCATGTCGCTGAGGCAGAAATTGAAACTGGCATTGCAACCAAAGAAGGAGGAAAAAAC TGCTGCAGGTCTCGGAGGAATTAGTCATGGAGGCCAAGGCTGCTTTTGAGGATGCTCAGGAGGAAGCCAGAGCGG AGAAGCTCCGAGAAGCACTTCCACCATTAGTGGCAGACAAAGGCATCGAGGCAGCCGCAGAAGTTGTCTGCGAAG CAAATGACCGTATGATCGGACAGTATATCGTTGTCTCGCCAAACTCTGTGCTGAAGAATGCCAAACTCGCACCAG CGCACCCGCTAGCAGATCAGGTTAAGATCATAACACACTCCGGAAGATCAGGAAGGTACGCGGTCGAACCATACG AAGAGGAGCAGTACAAGGTTACAAAGGCAGAGCTTGCAGAAACAGAGTACGTGTTTGACGTGGACAAGAAGCGTT TGGAGGGACTGAAGACCCGACCTGCGGTCCCGTACAAGGTCGAAACAATAGGAGTGATAGGCACACCGGGGTCGG **AAATTGAGGCCGACGTGCTAAGACTGAGGGGTATGCAGATTACGTCGAAGACAGTAGATTCGGTTATGCTCAACG** GATGCCACAAAGCCGTAGAAGTGCTGTACGTTGACGAAGCGTTCGCGTGCCACGCAGGAGCACTACTTGCCTTGA TTGCTATCGTCAGGCCCCGCAAGAAGGTAGTACTATGCGGAGACCCCATGCAATGCGGATTCTTCAACATGATGC AACTAAAGGTACATTTCAATCACCCTGAAAAAGACATATGCACCAAGACATTCTACAAGTATATCTCCCGGCGTT GCACACAGCCAGTTACAGCTATTGTATCGACACTGCATTACGATGGAAAGATGAAAACCACGAACCCGTGCAAGA AGAACATTGAAATCGATATTACAGGGGCCACAAAGCCGAAGCCAGGGGATATCATCCTGACATGTTTCCGCGGGT GGGTTAAGCAATTGCAAATCGACTATCCCGGACATGAAGTAATGACAGCCGCGGCCTCACAAGGGCTAACCAGAA **AAGGAGTGTATGCCGTCCGGCAAAAAGTCAATGAAAACCCACTGTACGCGATCACATCAGAGCATGTGAACGTGT** TGCTCACCCGCACTGAGGACAGGCTAGTGTGGAAAACCTTGCAGGGCGACCCATGGATTAAGCAGCTCACTAACA TACCTAAAGGAAACTTTCAGGCTACTATAGAGGACTGGGAAGCTGAACACAAGGGAATAATTGCTGCAATAAACA GCCCACTCCCCGTGCCAATCCGTTCAGCTGCAAGACCAACGTTTGCTGGGCGAAAGCATTGGAACCGATACTAG CCACGGCCGGTATCGTACTTACCGGTTGCCAGTGGAGCGAACTGTTCCCACAGTTTGCGGATGACAAACCACATT AGAGCATCCCACTAACGTACCATCCCGCCGATTCAGCGAGGCCGGTAGCTCATTGGGACAACAGCCCAGGAACCC GCACACAACTTGATTTGCAGACGGGGAGAACCAGAGTTATCTCTGCACAGCATAACCTGGTCCCGGTGAACCGCA ATCTTCCTCACGCCTTAGTCCCCGAGTACAAGGAGAAGCAACCCGGCCCGGTCGAAAAATTCTTGAACCAGTTCA AACACCACTCAGTACTTGTGGTATCAGAGGAAAAAATTGAAGCTCCCCGTAAGAGAATCGAATGGATCGCCCCGA TTGGCATAGCCGGTGCAGATAAGAACTACAACCTGGCTTTCGGGTTTCCGCCGCAGGCACGGTACGACCTGGTGT TCATCAACATTGGAACTAAATACAGAAACCACCACTTTCAGCAGTGCGAAGACCATGCGGCGACCTTAAAAACCC TTTCGCGTTCGGCCCTGAATTGCCTTAACCCAGGAGGCACCCTCGTGGTGAAGTCCTATGGCTACGCCGACCGCA ACAGTGAGGACGTAGTCACCGCTCTTGCCAGAAAGTTTGTCAGGGTGTCTGCAGCGAGACCAGATTGTGTCTCAA

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GCCGTGCCATCTATAAACGTTGGCCGACCAGTTTTACCGATTCAGCCACGGAGACAGGCACCGCAAGAATGACTG TGTGCCTAGGAAAGAAGTGATCCACGCGGTCGGCCCTGATTTCCGGAAGCACCCAGAAGCAGAAGCCTTGAAAT TGCTACAAAACGCCTACCATGCAGTGGCAGACTTAGTAAATGAACATAACATCAAGTCTGTCGCCATTCCACTGC TATCTACAGGCATTTACGCAGCCGGAAAAGACCGCCTTGAAGTATCACTTAACTGCTTGACAACCGCGCTAGACA GCTTGAAGGGAAGAAAGGGATTCAGTACTACAAAAGGAAAATTGTATTCGTACTTCGAAGGCACCAAATTCCATC AAGCAGCAAAAGACATGGCGGAGATAAAGGTCCTGTTCCCTAATGACCAGGAAAGTAATGAACAACTGTGTGCCT AAACGTTGCCGTGCCTTTGCATGTATGCCATGACGCCAGAAAGGGTCCACAGACTTAGAAGCAATAACGTCAAAG AAGTTACAGTATGCTCCTCCACCCCCCTTCCTAAGCACAAAATTAAGAATGTTCAGAAGGTTCAGTGCACGAAAG CTCCTCCTGCACAGGCCGAGGGGCCCCCGAAGTTGTAGCGACACCGTCACCATCTACAGCTGATAACACCTCGC TTGATGTCACAGACATCTCACTGGATATGGATGACAGTAGCGAAGGCTCACTTTTTTCGAGCTTTAGCGGATCGG ACAACTCTATTACTAGTATGGACAGTTGGTCGTCAGGACCTAGTTCACTAGAGATAGTAGACCGAAGGCAGGTGG TGGTGGCTGACGTTCATGCCGTCCAAGAGCCTGCCCCTATTCCACCGCCAAGGCTAAAGAAGATGGCCCGCCTGG ${\tt CAGCGGCTAGAAAAGAGCCCACTCCACCGGCAAGCAATAGCTCTGAGTCCCTCCACCTCTTTTTGGTGGGGTAT}$ CCATGTCCCTCGGATCAATTTTCGACGGAGAGACGGCCCGCCAGGCAGCGGTACAACCCCTGGCAACAGGCCCCA CGGATGTGCCTATGTCTTTCGGATCGTTTTCCGACGGAGAGATTGATGAGCTGAGCCGCAGAGTAACTGAGTCCG **AACCCGTCTGTTTGGATCATTTGAACCGGGCGAAGTGAACTCAATTATATCGTCCCGATCAGCCGTATCTTTTC** CACTACGCAAGCAGACGTAGACGCAGGAGCAGGAGGACTGAATACTGACTAACCGGGGTAGGTGGGTACATAT TTTCGACGGACACAGGCCCTGGGCACTTGCAAAAGAAGTCCGTTCTGCAGAACCAGCTTACAGAACCGACCTTGG AGCGCAATGTCCTGGAAAGAATTCATGCCCCGGTGCTCGACACGTCGAAAGAGAGAACAACTCAAACTCAGGTACC AGATGATGCCCACCGAAGCCAACAAAAGTAGGTACCAGTCTCGTAAAGTAGAAAATCAGAAAGCCATAACCACTG AGCGACTACTGTCAGGACTACGACTGTATAACTCTGCCACAGATCAGCCAGAATGCTATAAGATCACCTATCCGA TGCATGAGAACTATCCGACAGTAGCATCTTATCAGATTACTGACGAGTACGATGCTTACTTGGATATGGTAGACG GGACAGTCGCCTGCCTGGATACTGCAACCTTCTGCCCCGCTAAGCTTAGAAGTTACCCGAAAAAACATGAGTATA GAGCCCCGAATATCCGCAGTGCGGTTCCATCAGCGATGCAGAACACGCTACAAAATGTGCTCATTGCCGCAACTA AAAGAAATTGCAACGTCACGCAGATGCGTGAACTGCCAACACTGGACTCAGCGACATTCAATGTCGAATGCTTTC GAAAATATGCATGTAATGACGAGTATTGGGAGGAGTTCGCTCGGAAGCCAATTAGGATTACCACTGAGTTTGTCA CCGCATATGTAGCTAGACTGAAAGGCCCTAAGGCCGCCCCCATTTTGCAAAGACGTATAATTTGGTCCCATTGC AATTAGTGCGTAGGCTTACGGCCGTCTTGCTTCCAAACATTCACACGCTTTTTGACATGTCGGCGGAGGATTTTG ATGCAATCATAGCAGAACACTTCAAGCAAGGCGACCCGGTACTGGAGACGGATATCGCATCATTCGACAAAAGCC **AATCCGGAATGTTCCTCACACTTTTTGTCAACACAGTTTTGAATGTCGTTATCGCCAGCAGAGAGTACTAGAAGAGC** GGCTTAAAACGTCCAGATGTGCAGCGTTCATTGGCGACGACAACATCATACATGGAGTAGTATCTGACAAAGAAA TGGCTGAGAGGTGCGCCACCTGGCTCAACATGGAGGTTAAGATCATCGACGCAGTCATCGGTGAGAGACCACCTT ACTTCTGCGGCGGATTTATCTTGCAAGATTCGGTTACTTCCACAGCGTGCCGCGTGGCGGACCCcctgaaaaggc tgtttaagttgggtaaaccgctcccagccgacgacgagcaagacgagacagaagacgcgctctgctagatgaaa caaaggcgtggtttagagtaggtataacaggcactttagcagtggccgtgacgacccggtatgaggtagacaata ttacacctgtcctactggcattgagaacttttgcccagagcaaaagagcattccaagccatcagaggggaaataa agcatctctacggtggtcctaaatagtcagcatagtacatttcatctgactaatactacaacaccaccatga ${\tt atagaggattctttaacatgctcggccgccgccccttcccggccccactgccatgtggaggccgcggAGAAGGA}$ GGCAGGCGCCCCGATGCCTGCCCGCAACGGGCTGGCTTCTCAAATCCAGCAACTGACCACAGCCGTCAGTGCCC TAGTCATTGGACAGGCAACTAGACCTCAACCCCCACGTCCACGCCCACCCGCCACAGAAGAAGCAGGCGCCCA AGAGACAGCGCATGGCACTTAAGTTGGAGGCCGACAGATCGTTCGACGTCAAGAACGAGGACGGAGATGTCATCG GGCACGCACTGGCCATGGAAGGAAAGGTAATGAAACCTCTGCACGTGAAAGGAACCATCGACCACCCTGTGCTAT CAAAGCTCAAATTTACCAAGTCGTCAGCATACGACATGGAGTTCGCACAGTTGCCAGTCAACATGAGAAGTGAGG CATTCACCTACACCAGTGAACACCCCGAAGGATTCTATAACTGGCACCACGGAGCGGTGCAGTATAGTGGAGGTA GATTTACCATCCCTCGCGGAGTAGGAGGCAGAGGAGACAGCGGTCGTCCGATCATGGATAACTCCGGTCGGGTTG





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TCGCGATAGTCCTCGGTGGAGCTGATGAAGGAACACGAACTGCCCTTTCGGTCGTCACCTGGAATAGTAAAGGGA AGACAATTAAGACGACCCCGGAAGGGACAGAAGAGTGGTCCGCAGCACCACTGGTCACGGCAATGTGTTTGCTCG GAAATGTGAGCTTCCCATGCGACCGCCCGCCCACATGCTATACCCGCGAACCTTCCAGAGCCCTCGACATCCTTG AAGAGAACGTGAACCATGAGGCCTACGATACCCTGCTCAATGCCATATTGCGGTGCGGATCGTCTGGCAGAAGCA AAAGAAGCGTCACTGACGACTTTACCCTGACCAGCCCCTACTTGGGCACATGCTCGTACTGCCACCATACTGAAC CGTGCTTCAGCCCTGTTAAGATCGAGCAGGTCTGGGACGAAGCGGACGATAACACCATACGCATACAGACTTCCG CCCAGTTTGGATACGACCAAAGCGGAGCAGCAAGCGCAAACAAGTACCGCTACATGTCGCTTAAGCAGGATCACA CCGTTAAAGAAGGCACCATGGATGACATCAAGATTAGCACCTCAGGACCGTGTAGAAGGCTTAGCTACAAAGGAT ACTTTCTCCTCGCAAAATGCCCTCCAGGGGACAGCGTAACGGTTAGCATAGTGAGTAGCAACTCAGCAACGTCAT GTACACTGGCCCGCAAGATAAAACCAAAATTCGTGGGACGGGAAAAATATGATCTACCTCCCGTTCACGGTAAAA AAATTCCTTGCACAGTGTACGACCGTCTGAAAGGAACAACTGCAGGCTACATCACTATGCACAGGCCGGGACCGC ACGCTTATACATCCTACCTGGAAGAATCATCAGGGAAAGTTTACGCAAAGCCGCCATCTGGGAAGAACATTACGT ATGAGTGCAAGTGCGGCGACTACAAGACCAGAACCGTTTCGACCCGCACCGAAATCACTGGTTGCACCGCCATCA AGCAGTGCGTCGCCTATAAGAGCGACCAAACGAAGTGGGTCTTCAACTCACCGGACTTGATCAGACATGACGACC ACACGCCCAAGGGAAATTGCATTTGCCTTTCAAGTTGATCCCGAGTACCTGCATGGTCCCTGTTGCCCACGCGC CGAÁTGTAATACATGGCTTTAAACACATCAGCCTCCAATTAGATACAGACCACTTGACATTGCTCACCACCAGGA GACTAGGGGCAAACCCGGAACCACCACTGAATGGATCGTCGGAAAGACGGTCAGAAACTTCACCGTCGACCGAG ATGGCCTGGAATACATATGGGGAAATCATGAGCCAGTGAGGGTCTATGCCCAAGAGTCAGCACCAGGAGACCCTC **ACGGATGGCCACACGAAATAGTACAGCATTACTACCATCGCCATCCTGTGTACACCATCTTAGCCGTCGCATCAG** CATACGCCCTGGCCCCAAACGCCGTAATCCCAACTTCGCTGGCACTCTTGTGCTGCGTTAGGTCGGCCAATGCTG AAACGTTCACCGAGACCATGAGTTACTTGTGGTCGAACAGTCAGCCGTTCTTCTGGGTCCAGTTGTGCATACCTT CGAAGGTAGACGCCTACGAACATGCGACCACTGTTCCAAATGTGCCACAGATACCGTATAAGGCACTTGTTGAAA ACATTACCTGCAAATTCACCACTGTGGTCCCCTCCCCAAAAATCAAATGCTGCGGCTCCTTGGAATGTCAGCCGG CCGTTCATGCAGACTATACCTGCAAGGTCTTCGGAGGGGTCTACCCCTTTATGTGGGGAGGAGCGCAATGTTTTT GCGACAGTGAGAACAGCCAGATGAGTGAGGCGTACGTCGAACTGTCAGCAGATTGCGCGTCTGACCACGCGCAGG CGATTAAGGTGCACACTGCCGCGATGAAAGTAGGACTGCGTATAGTGTACGGGAACACTACCAGTTTCCTAGATG TGTACGTGAACGGAGTCACACCAGGAACGTCTAAAGACTTGAAAGTCATAGCTGGACCAATTTCAGCATCGTTTA CGCCATTCGATCATAAGGTCGTTATCCATCGCGGCCTGGTGTACAACTATGACTTCCCGGAATATGGAGCGATGA AACCAGGAGCGTTTGGAGACATTCAAGCTACCTCCTTGACTAGCAAGGATCTCATCGCCAGCACAGACATTAGGC TACTCAAGCCTTCCGCCAAGAACGTGCATGTCCCGTACACGCAGGCCGCATCAGGATTTGAGATGTGGAAAAACA ACTCAGGCCGCCCACTGCAGGAAACCGCACCTTTCGGGTGTAAGATTGCAGTAAATCCGCTCCGAGCGGTGGACT GTTCATACGGGAACATTCCCATTTCTATTGACATCCCGAACGCTGCCTTTATCAGGACATCAGATGCACCACTGG TCTCAACAGTCAAATGTGAAGTCAGTGAGTGCACTTATTCAGCAGACTTCGGCGGGATGGCCACCCTGCAGTATG TATCCGACCGCGAAGGTCAATGCCCCGTACATTCGCATTCGAGCACAGCAACTCTCCAAGAGTCGACAGTACATG TCCTGGAGAAAGGAGCGGTGACAGTACACTTTAGCACCGCGAGTCCACAGGCGAACTTTATCGTATCGCTGTGTG GGAAGAAGACACATGCAATGCAGAATGTAAACCACCAGCTGACCATATCGTGAGCACCCCGCACAAAAATGACC **AAGAATTTCAAGCCGCCATCTCAAAAACATCATGGAGTTGGCTGTTTGCCCTTTTCGGCGGCGCCCTCGTCGCTAT TAATTATAGGACTTATGATTTTTGCTTGCAGCATGATGCTGACTAGCACGAAGATGACCGCTACGCCCCAATG** ATCCGACCAGCAAAACTCGATGTACTTCCGAGGAACTGATGTGCATAATGCATcaggctggtacattagatcccc gcttaccgcgggcaatatagcaacactaaaaactcgatgtacttccgaggaagcgcagtgcataatgctgcgcag tgttgccacataaccactatattaaccatttatctagcggacgccaaaaactcaatgtatttctgaggaagcgtg gtgcataatgccacgcagcgtctgcataacttttattatttcttttattaatcaacaaaattttgtttttaacat ttc

FIG. 2C-3

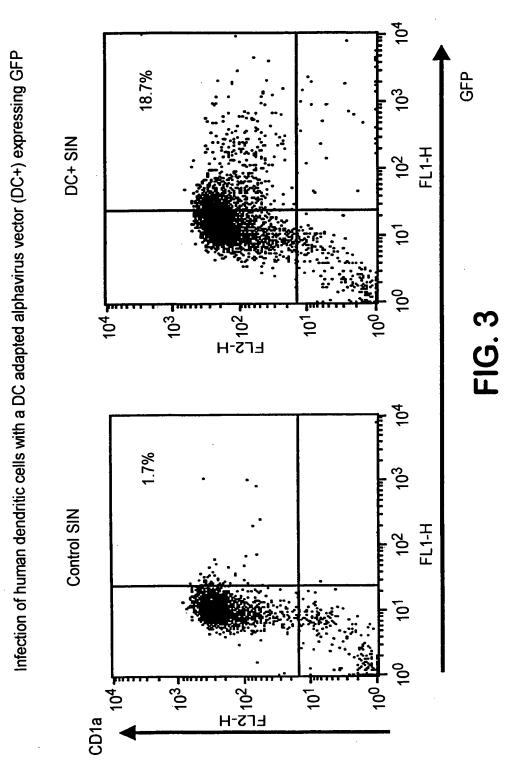


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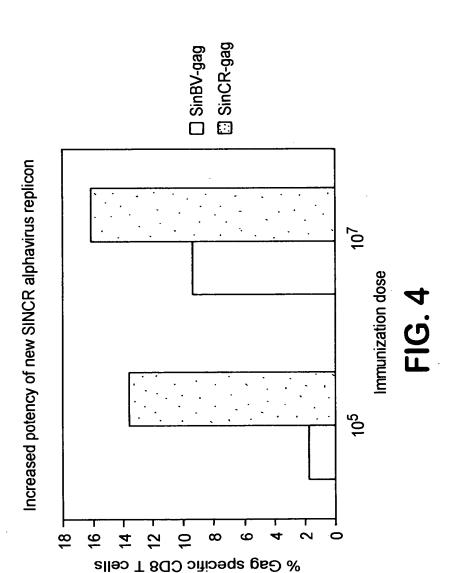
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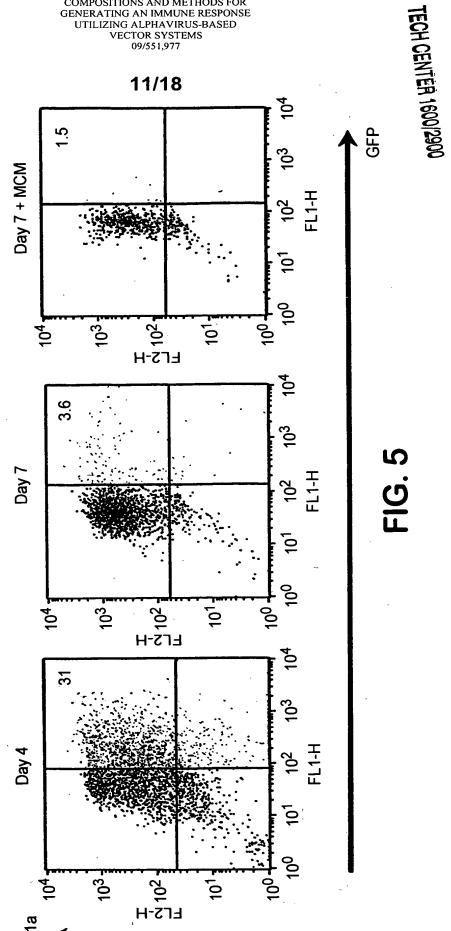


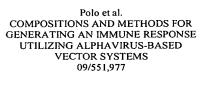
ERY & TRADE

DC+ SIN vectors target immature human dendritic cells

immature

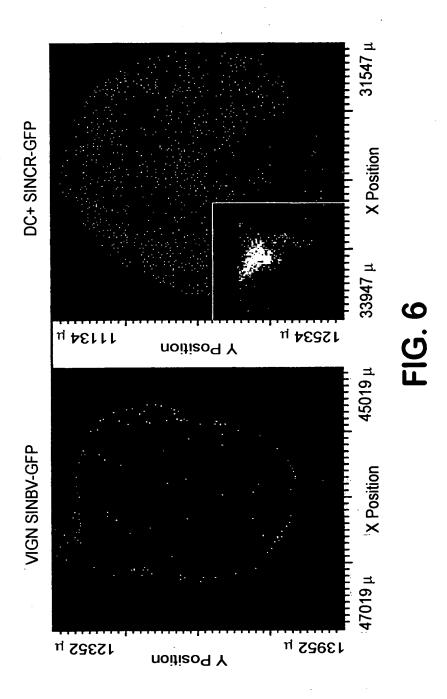
Polo et al. COMPOSITIONS AND METHODS FOR GENERATING AN IMMUNE RESPONSE UTILIZING ALPHAVIRUS-BASED VECTOR SYSTEMS 09/551,977





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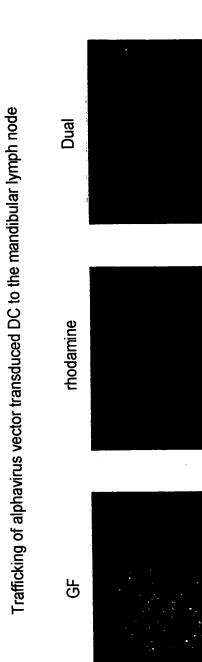












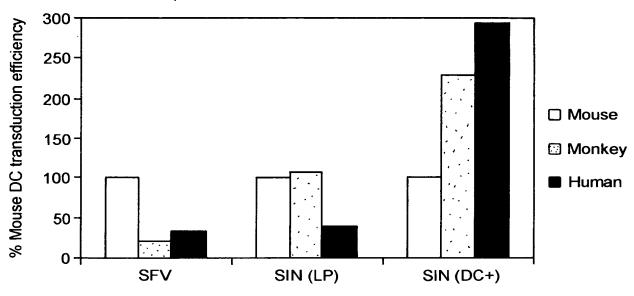
SIN-GFP vector injected intradermally, with rhodamine paint applied to skin

x20



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Mouse DC transduction is not predictive of the ability of alphavirus vectors to transduce human DC

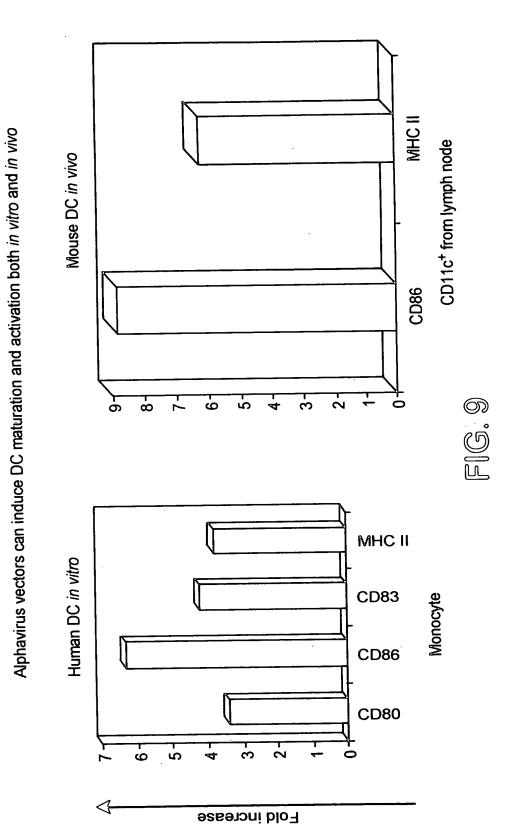


Source of structural proteins

FIG. 8



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8

8

2

9

20

8

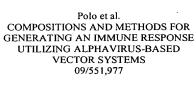
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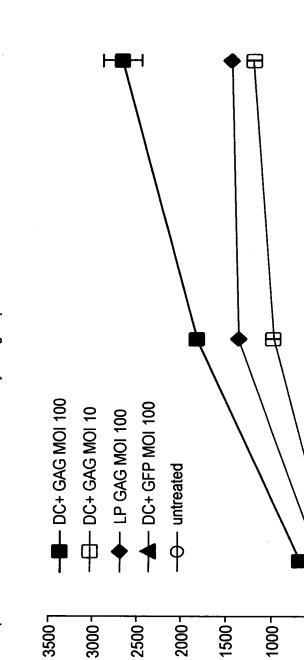
500-

CD11c+ DC (x 10³)/well

FIG. 10



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(lm/gq) 2-JI

Adapted alphavirus vectors can be used to assay antigen presentation and immune stimulation in vitro

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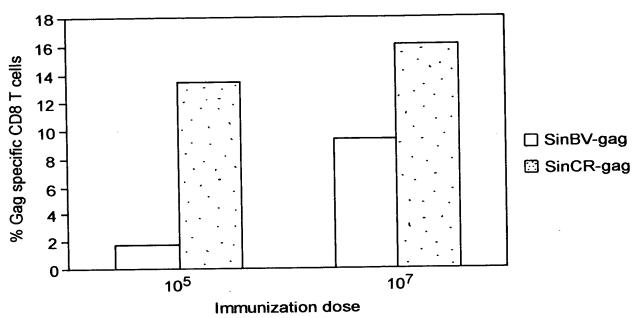
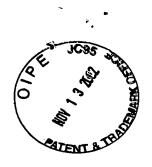


FIG. 11



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Enhanced immune response by using a prime-boost strategy

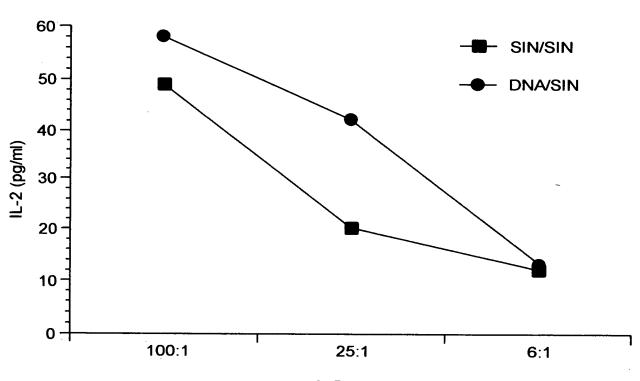


FIG. 12